## **Description of Additional Supplementary Files**

File Name: Supplementary Movie 1

Description: Wild-type *C. difficile* spores internalize in small intestine. Travel through a confocal *Z*-stack from the apical face of the small intestine. F-actin is visualized with fluorescently labeled phalloidin with Alexa-Fluor 488 (green) to distinguish the structure of the intestinal villus (green). And *C. difficile* spores were detected using anti-*C. difficile* chicken spore antibody IgY and goat anti-chicken IgY secondary antibody conjugated with Alexa-Fluor 568 (red). DNA was stained with DAPI (blue). Arrows indicate internalized spores.

File Name: Supplementary Movie 2

Description: Wild-type *C. difficile* spores internalize in colonic mucosa. Travel through a confocal *Z*-stack from the apical face of the colonic mucosa. F-actin is visualized with fluorescently labeled phalloidin Alexa-Fluor 488 (green) to distinguish the structure of the colonic crypt, and *C. difficile* spores were detected using anti-*C. difficile* chicken spore antibody IgY and goat anti-chicken IgY secondary antibody conjugated with Alexa-Fluor 568 (red). Arrows indicate internalized spores.